

REMARKS

Enclosed please find Figure 1 consisting of 1 (one) sheet marked in red to identify the changes thereto. A discussion describing the revisions and the support therefore in the originally filed disclosure is provided below. Also enclosed please find Figure 1 consisting of 1 (one) sheet of the formal drawing for the subject application. In accordance with 37 C.F.R. 1.84(c), identifying indicia are provided on the backside of the sheet.

Applicants respectfully request that, prior to examination, Figure 1 be amended as shown on the marked-up version of Figure 1 and that pages 1 and 2 of the sequence listing be substituted with pages 1 and 2 submitted herewith. Applicants respectfully submit that no new matter is being added by the amendment of this figure and substitute sequence listing.

Figure 1 was originally filed with 9 (nine) residues inadvertently deleted from each row on the right hand side of the figure as indicated in the marked-up version submitted herewith. New Figure 1 sets forth the complete sequence for each organism. Applicants respectfully submit that, with the exception of the *Drosophila* ("fly") sequence, all the sequences in Figure 1 are known in the prior art, as described with genome database citations on page 3, lines 11-14 of the specification. Thus, one of ordinary skill in the pertinent art would have been able to obtain the correct sequences. In addition, the inadvertently omitted residues from the *Drosophila* sequence are described in the originally filed sequence listing in Sequence I.D No.1. Thus, no new matter has been added by virtue of this amendment.

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Applicants further request that pages 1 and 2 of the sequence listing be replaced with the substituted pages submitted herewith. The Sequence I.D. No.1 in the original sequence listing was filed with 1 residue (K) inadvertently omitted in the second line of the listing (EIKSLED in the original should read EIKKSLED). This portion of the sequence was given correctly in the second line of the *Drosophila* ("fly") sequence in Figure 1 as originally filed and was also correctly shown in Figure 6B as originally filed. In addition, Sequence I.D. No.2 in the original sequence listing shows the DNA sequence encoding the correct protein, enabling one of ordinary skill in the pertinent art to obtain the correct sequence. Therefore, Applicants respectfully submit that no new matter is being added by the amendment of this sequence listing.

Should the Examiner wish to discuss the above amendment, the undersigned attorney would appreciate the opportunity to do so.

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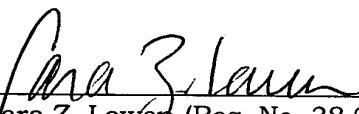
S. Cohen, et al.
U.S.S.N. Not Assigned
Preliminary Amendment
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Applicants believe that additional fees are not required for consideration of the within Preliminary Amendment. However, if for any reason a fee is required, a fee paid is inadequate or credit is owed for any excess fee paid, you are hereby authorized and requested to charge Deposit Account No. **04-1105**.

Respectfully submitted,

Date: 5/2/01

By:


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Figure 1:

multiple sequence alignment

yeast	MSTLIPPPSKQQKKEAQLPREVAAIPKDLNVSIFQALDTGDNVGGALRVPAGISEKQL	
c.elegans	-----PQISVSEDENELG---GSGILVPVDISTNEL	
fly	-----MQETDTEQEATPHTIQARLVYTGEAGPPIDLPAGITTQQL	
mouse	-----EEAAAGD-VQRLVQFQDEGGQLLGSPFDVPVDITPDKL	
human	-----GSPFDVPVDITPDRL	
frog	-----MKEDVGRLLIQFKNENGEGLGPFDVPLDITPDKL	
	* : : * : ..*	
yeast	EELLNQLNGTSDDPVPYTFSCTIQGKKASDPVKTIDITDNLYSSLIKPGYNSTEDQITLL	
c.elegans	QILCNQLLGSR---FCLNNEFSVSG-----AEIVDHSIRKSLEEIDFET---LKV	
fly	GLICCNALLKNE---EATPYLFFVGE-----DEIKKSLEDTLDSLAVDT-ENVIDIV	
mouse	XLVCNALL-AQEEPLPLAFVHD-----AEIVVSSLGKTLESQSV-ETEKIVDII	
human	QLVCNALL-AQEDPCPLAFVHD-----AEIVVSSLGKTLESQAV-ETEKVLDIY	
frog	QLVCNALL-QEEDPVPLAFVQD-----LEIVTSLDKTLEKQSV-ETEKVIDII	
	* : .. : * : : : :	
yeast	YT普RAVFKVVKPVRTSSAIAGHGSTIILCSAFAPHTSSRMVTGAGDNTARIWDCDTQTPMH	
c.elegans	YQPQAVFVRVPVRTCSASIPGHGEPVISAQFSPDGRC-LASGSGDQTMRWIELELPLH	
fly	YQPQAVFKVPRVTRCTSSMPGHAEEAVVSLNFSPDGAH-LASGSGDTTVRLWDLNTETPHF	
mouse	YQPQAVFVRVRAVTRCTS-----	
human	YQPQAVFKVRAVTRCTSSLEGTEAVISVAFSPTGKY-LASGSGDTTVRFWDLSTETPHF	
frog	* : : * : : * : : :	
yeast	TLKGHYNWVLCVSWSPDGEVIATGSMNTIRLWDPKSGQCLGDAKGSKWITSLSWEP	
c.elegans	TCKSHKSWVLCIAWSPDATKIASACKNGEICIWNAKTGEQIGKTLKRHKQWIXXLAWQP-	
fly	TCTGHKQWVLCVSWAPDGKRLASGCKAGSIIWDPETGQQKGRPLSGHKHHINCLAWEPY	
mouse	-----	
human	-----	
frog	TSKGHTHWVLSIAWSPDGKKLASGCKNSQIFIWDPSTGKQIGKPLTGHSKWITWLCWEPL	
yeast	LVKPGSKPRLASSSKDGTIKIWDTSRVCQYTMMSGHTNSVSCVKWGGQGLIYSGSHDRTV	
c.elegans	-----TVKMWR-----ADDGVMCRNMTG-----	
fly	HRDPECR-KLASASGDGDRCIWDVKGQCLMNIAGHTNAVTAVRWGGAGLIYTSSKDRTV	
mouse	-----	
human	-----	
frog	HLNPESTRY-LASASKDCTIRIWDVTMVGQCQKILTSHTQSVTAVKWGGDGLIYSSSQDRTI	
yeast	RVWDINSQGRCINILKSHAHWVNHLSSLSTDYALRIGAFDHTGKK-----PSTPEEAQKKA	
c.elegans	-----HAHWINTLALNTDYALRTSCFE-----PSK-----	
fly	KMWR-AADGILCRTFSGHAWVNNIALSTDYVLRTGPFHVKDRSKSHLSI STEELQESA	
mouse	-----	
human	-----	
frog	KAWR-AQDGVLCTRLQGHAHWVNTMALSTDYVLRKGAFNPADAS--VNPQLMSGSLEVLK	
yeast	LENYEKICKKNGNSEEMMVTAASDDYTMFLWNPLKSTKPIARMTHQKLVNHVAFSPDGRY	
c.elegans	-----INRMTGHMQLVNCVVFSPDTRY-----	
fly	LKRYQAVCP---DEVESLVCSDDNLYLWRN-NQNKCVERMTGHQNVVNDVKYSPDVKL	
mouse	-----	
human	-----	
frog	EKALKRSNEVRGQGPERLVSGSEDFTLFLWAPAEKKPLQRMTHQALINEVLFSPDTRI	
yeast	IVSASFDSIKLWDGRDGKFISTFRGHIASVYQVAASSDCRLLVSCSKDTTLKVWDVRTR	
c.elegans	IASASFDSKVKLWCGRTGKYLASFGRHVGPVYQVAWSADSRLLVSGSADSTLKVFEKTK	
fly	IASASFDSKVKLWRASDGQYMATFRGHVQAVYTVAWSADSRLIVSGSKDSTLKVWSVQTK	
mouse	-----	
human	-----	
frog	IASASFDSIKLWDGKTGKFLTSRGHVSAYQIAWSADSRLLVSGSSDSTLKVWDSKTK	
yeast	KLSVDLPGIJKTKLY-VDWSVDGKRVCSGGKDKMVRLWTH	
c.elegans	SLYYDLPGHGDEVFTWDWSPEGTKVVSGGKDKVLIKWL--	
fly	KLAQELPGHADEVFGVDWAPDGSRVASGGKDKVLIK WAY	
mouse	-----	
human	-----	
frog	KLLIDLPGHADEVYSVDWSPDGQRVASGGKDKCLRIWRK	

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Sequence I.D. No.1: Drosophila Notchless protein

MQETDTEQEATPHTIQARLVYTGEAGPPIDLPAGITTQQLGLICNALLKNEEA
TPYLFFVGEDEIKSLEDTLASVDTENVIDIVYQPQAVFKVRPVTRCTSSMP
5 GHAEAVVSLNFSPDGAHLASGSGDTTVRLWDLNTETPHFTCTGHKQWVLCV
SWAPDGKRLASGCKAGSIIWDPETGQQKGRPLSGHKKHINCLAWEPYHRDP
ECRKLASASGDGDCRIWDVKGQCLMNIAGHTNAVTAVRWGGAGLIYTSSK
DRTVKMWRAADGILCRTFSGHAHWVNNIALSTDYVLRTGPFHPVKDRSKSH
LSLSTEELQESALKRYQAVCPDEVESLVSCSDNTLYLWRNNQNKCVERMT
10 GHQNVVNDVKYSPDVKLIASASFDSVRLWRASDGQYMATFRGHVQAVYT
VAWSADSRLIVSGSKDSTLVWSVQTKLAQELPGHADEVFGVDWAPDGSR
VASGGKDKVIKLWAY

Sequence I.D. No. 2: Drosophila *Nle* cDNA

15 aattcccaaaaaATGCAGGAGACGGACACGGAGCAAGAGGCCACGCCACATACG
ATACAGGCAGCGCCTCGTTACACGGCGAGGAAGCCGGCCGCCAATCGA
CCTGCCGGCAGGAATCACTACCCAGCAATTGGACTGATTGCAACGCGC
TGCTGAAAAACGAGGAAGCCACTCCATATTGTTTCTGGCGAGGAT
20 GAGATCAAGAACGAGCCTGGAGGACACGTTGGACTTGGCTCAGTGGACA
CCGAAAACGTGATCGATTGTGTATCAGCCACAGGCCTTCAAAGTG
CGCCCAGTGACAAGATGCACGAGTTCCATGCCGGACACGCCGAGGCTGT
GGTTTCGCTGAATTTCAGCCGGATGGTGTCTCATCTGCCAGTGGAAAGTG
GCGACACCACAGTGCATTGTGGATCTAACACAGAGACACCGCACTTC
25 ACCTGCACAGGTATAAGCAGTGGTTCTGTGCGTATCCTGGCTCCGGA
TGGCAAACGGTTGCCAGCGGTTGCAAAGCGGGCTCTATAATCATCTGGG
ACCCGGAGACGGGTCAAGCAGAAGGGCGACCCCTGAGTGGCACAAGAA
ACACATCAACTGCCCGCTGGAACCGTATCGCGATCCGGAGTGCA
GGAAACTTGCTCCGCCAGTGGAGACGGGACTGCCGGATTGGACGTA
30 AAATTGGGCCAGTGCCTTATGAACATTGCCGGACACACAAATGCTGTGAC
AGCAGTGAAGATGTGGCGAGCAGCTGATGGAATCTTGTGCCGGACGTT
GCACAGTGAAGATGTGGCGAGCAGCTGATGGAATCTTGTGCCGGACGTT
TCTGGCCAAGCTCACTGGGTAAACAAACATTGCGCTGAGCACCGATTACGT
CCTGCGCACTGGTCCATTCCATCCGGTAAGGATCGCTCCAAGAGCCACC

TCAGTTGAGCACTGAGGAATTGCAGGAATCTGCCTGAAGCGCTACCAAG
GCCGTGTGCCCTGACGAGGTGGAGTCGCTGGTTCTGTTGGATGACAA
CACCCCTCTATCTGTGGCGGAACAACCAGAACAGAAGTGCCTGAGCGCATGA
CAGGGCACCAGAACGTGGTCAACGATGTGAAATATTGCCGGATGTAAAG
5 CTAATTGCGTCTGCTTCATTGACAAGTCAGTGCCTGTGGCGAGCCAGC
GATGGTCAGTACATGGCCACCTTCCGGGTATGTGCAGGCTGTTACAC
GGTTGCCTGGTCCCGGGACTCCCGCTTGATTGTTCCGGCAGCAAAGACTC
AACTCTAAAAGTATGGAGTGTGCAGACGAAGAAACTGGCACAGGAGCTG
CCTGGACATGCGGATGAGGTGTTGGAGTGGACTGGCGCCGATGGCTC
10 TAGAGTTGCCTCTGGTGGCAAGGACAAAGTTATAAGCTATGGGCTTATT
AAc aaatcattaacttgtacacggtaaagaaaacttaggaataaagtaaaacgtcctgagtaaaaaaaaaaaaaaaaaaaaaaa
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